

Listing of Claims

Please amend claims 1, 4, 29, 34-36, 38 and 41, and please add new claims 43-48, as shown below. This listing of claims will replace all prior versions and listings of claims in the instant application.

1. (Currently Amended) An antimicrobial sulfonamide derivative, or a salt or a hydrate thereof, comprising:

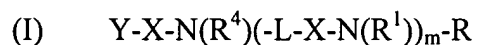
a core cyclic peptide or core antibiotic of ~~a~~ an acidic lipopeptide antibiotic; and
a lipophilic moiety,

wherein said lipophilic moiety is covalently attached to the core cyclic peptide or core antibiotic *via* a linking chain which includes a sulfonamide linkage and wherein said core cyclic peptide or core antibiotic is not of laspartomycin ~~or polymyxin~~.

2. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a sulfonamide linkage.

3. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a linker that links the core cyclic peptide or core antibiotic to the lipophilic moiety.

4. (Currently Amended) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 which is a compound according to structural Formula (I):



wherein:

Y is a lipophilic moiety;

each X is independently selected from the group consisting of ~~eo-SO₂-~~
-CO-SO₂-, -CS-, -PO-, -OP(O)-, -OC(O)-, -NHCO and -N(R¹)CO- with the proviso that at least one X is ~~-SO₂-~~ -SO₂-;

M is 0 or 1;

L is a linker;

N is nitrogen;

R^1 and R^4 are each independently selected from the group consisting of hydrogen, (C_1-C_{25}) alkyl optionally substituted with one or more of the same or different R^2 groups, (C_1-C_{25}) heteroalkyl optionally substituted with one or more of the same or different R^2 groups, (C_5-C_{30}) aryl optionally substituted with one or more of the same or different R^2 groups, (C_5-C_{30}) arylaryl optionally substituted with one or more of the same or different R^2 groups, (C_5-C_{30}) biaryl optionally substituted with one or more of the same or different R^2 groups, five to thirty membered heteroaryl optionally substituted with one or more of the same or different R^2 groups, (C_6-C_{30}) arylalkyl optionally substituted with one or more of the same or different R^2 groups and six to thirty membered heteroarylalkyl optionally substituted with one or more of the same or different R^2 groups;

each R^2 is independently selected from the group consisting of $-OR^3$, $-SR^3$, $-NR^3R^3$, $-CN$, $-NO_2$, $-N^3-N_3$, $-C(O)OR^3$, $-C(O)NR^3R^3$, $-C(S)NR^3R^3$, $-C(NR^3)NR^3R^3$, $-CHO$, $-R^3CO$, $-SO_2R^3$, $-SOR^3$, $-PO(OR^3)_2$, $-PO(OR^3)$, $-CO_2H$, $-SO_3H$, $-PO_3H$, halogen and trihalomethyl;

each R^3 is independently selected from the group consisting of hydrogen, (C_1-C_6) alkyl, (C_5-C_{10}) aryl, five to sixteen membered heteroaryl, (C_6-C_{16}) arylalkyl and six to sixteen membered heteroarylalkyl; and

R is a core cyclic peptide or core antibiotic of ~~a~~ an acidic lipopeptide antibiotic, wherein said core cyclic peptide or core antibiotic is not of laspartomycin ~~or~~ polymyxin.

5. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of zaomycin, crystallomycin, aspartocin, amphomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimycin.

6. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of zaomycin, crystallomycin, aspartocin, amphomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimycin.

7. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145 or Antibiotic A-21978C.

8. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A54145 or Antibiotic A-21978C.

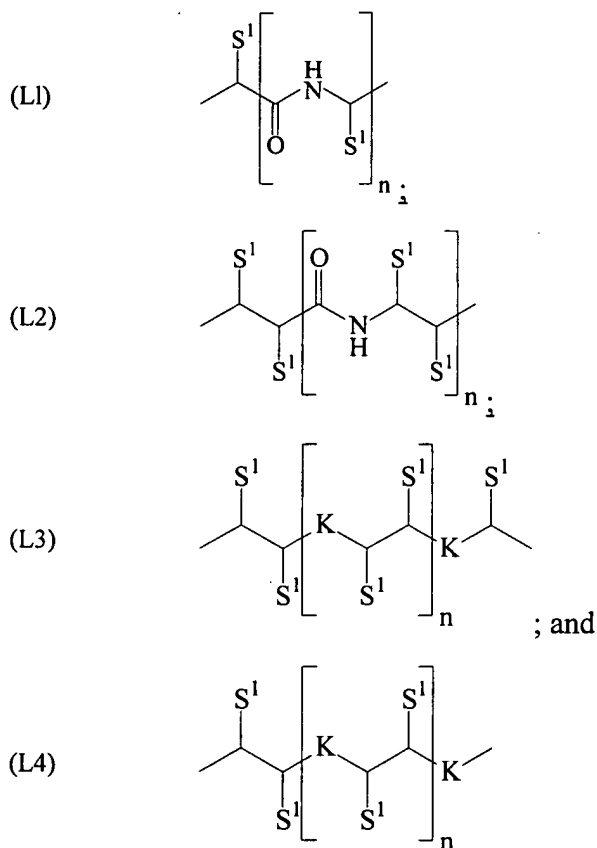
9. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of aspartocin.

10. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of aspartocin.

11. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which m is 1.

12. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which R¹ and R⁴ are hydrogen.

13. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 4 in which L is selected from the group consisting of:



or a pharmaceutically acceptable salt or hydrate thereof, wherein:

n is 0, 1, 2 or 3;

each S¹ is independently selected from the group consisting of hydrogen, (C₁-C₁₀) alkyl optionally substituted with one or more of the same or different R⁵ groups, (C₁-C₁₀) heteroalkyl optionally substituted with one or more of the same or different R⁵ groups, (C₅-C₁₀) aryl optionally substituted with one or more of the same or different R⁵ groups, (C₅-C₁₅) arylaryl optionally substituted with one or more of the same or different R⁵ groups, (C₅-C₁₅) biaryl optionally substituted with one or more of the same or different R⁵ groups, five to ten membered heteroaryl optionally substituted with one or more of the same or different R⁵ groups, (C₆-C₁₆) arylalkyl optionally substituted with one or more of the same or different R⁵ groups and six to sixteen membered heteroarylalkyl optionally substituted with one or more of the same or different R⁵ groups;

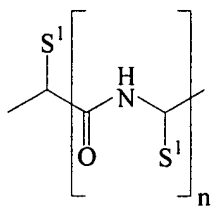
each R^5 is independently selected from the group consisting of $-OR^6$, $-SR^6$, $-NR^6R^6$, $-CN$, $-NO_2$, $-N_3$, $-C(O)OR^6$, $-C(O)NR^6R^6$, $-C(S)NR^6R^6$, $-C(NR^6)NR^6R^6$, $-CHO$, $-R^6CO$, $-SO_2R^6$, $-SOR^6$, $-PO(OR^6)_2$, $-PO(OR^6)$, $-CO_2H$, $-SO_3H$, $-PO_3H$, halogen and trihalomethyl;

each R^6 is independently selected from the group consisting of hydrogen, (C_1-C_6) alkyl, (C_5-C_{10}) aryl, five to sixteen membered heteroaryl, (C_6-C_{16}) arylalkyl and six to sixteen membered heteroarylalkyl; and

each K is independently selected from the group consisting of oxygen, nitrogen and sulfur.

14. (Original) The antimicrobial sulfonamide of Claim 13 in which each S^1 is independently a side-chain of a genetically encoded α -amino acid.

15. (Previously Presented) The antimicrobial sulfonamide of Claim 13 in which L is:



16. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which each S^1 is independently a side-chain of a genetically encoded α -amino acid.

17. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which n is 0.

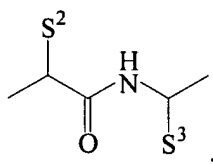
18. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 17 in which S^1 is hydrogen, Y is decan-yl and R is the core cyclic peptide of asparticin.

19. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S^1 is $-\text{CH}_2\text{-CO}_2\text{H}$, $-\text{CH}_2\text{-CH}_2\text{-CO}_2\text{H}$, $-\text{C(OH)H-CONH}_2$, $-\text{CH}_2\text{-CONH}_2$ or $-\text{CH}_2\text{-CH}_2\text{-CONH}_2$ or a salt or hydrate thereof.

20. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S^1 is $-\text{CH}_2\text{-indol-2-yl}$ or $-\text{CH}_2\text{-phenyl}$.

21. (Cancelled)

22. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



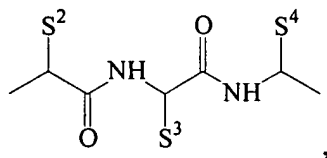
wherein S^2 and S^3 are each independently a side chain of a genetically encoded α -amino acid.

23. (Cancelled)

24. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S^2 is hydrogen, $-\text{CH}_2\text{-indol-2-yl}$, $-\text{CH}_2\text{-CONH}_2$ or $-\text{CH}_2\text{-CH}_2\text{-CONH}_2$ and S^3 is $-\text{CH}_2\text{-CO}_2\text{H}$, $-\text{CH}_2\text{-CH}_2\text{-CO}_2\text{H}$ or a salt or hydrate thereof.

25. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S^2 is $-\text{CH}_2\text{-CO}_2\text{H}$, $-\text{CH}_2\text{-CH}_2\text{-CO}_2\text{H}$ or a salt or hydrate thereof and S^3 is $-\text{C(OH)H-CONH}_2$.

26. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



wherein S^2 , S^3 , and S^4 are each independently a side chain of a genetically encoded α -amino acid.

27. (Cancelled)

28. (Original) The antimicrobial sulfonamide derivative of Claim 26 in which S^2 is $-\text{CH}_2\text{-indol-2-yl}$, S^3 is $-\text{CH}_2\text{-CONH}_2$ or $-\text{CH}_2\text{-CH}_2\text{-CONH}_2$ and S^4 is $-\text{CH}_2\text{-CO}_2\text{H}$, $-\text{CH}_2\text{-CH}_2\text{-CO}_2\text{H}$ or a salt or hydrate thereof.

29. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 26 in which S^2 is $-\text{CH}_2\text{-indol-2-yl}$, S^3 is $-\text{CH}_2\text{-CO}_2\text{H}$, $\text{CH}_2\text{-CH}_2\text{-CO}_2\text{H}$ or a salt or hydrate thereof and S^4 is $-\text{CH}_2\text{-CONH}_2$, $-\text{CH}_2\text{-CH}_2\text{-CONH}_2$ or $-\text{C(OH)H-CONH}_2$.

30. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which m is 0.

31. (Original) The antimicrobial sulfonamide derivative of Claim 30 in which R^4 is hydrogen.

32. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 30 in which R is the core antibiotic of aspartocin.

33. (Previously Presented) The antimicrobial sulfonamide derivative of Claim 30 in which R is the core cyclic peptide of aspartocin.

34. (Currently Amended) A pharmaceutical composition comprising an antimicrobial sulfonamide derivative according to ~~Claim 4~~ any one of Claims 1 to 5 and a pharmaceutically acceptable adjuvant, excipient, carrier or diluent.

35. (Currently Amended) A method for treating or preventing a microbial infection, said method comprising the step of administering to a subject a therapeutically effective amount of ~~a compound according to Claim 4 or a therapeutically effective amount of a~~ pharmaceutical composition according to Claim 34.

36. (Currently Amended) A method of inhibiting microbial growth, said method comprising the step of administering to a microbe an antimicrobially effective amount of ~~an antimicrobial sulfonamide derivative according to Claim 4 or an antimicrobially effective amount of a~~ pharmaceutical composition according to Claim 34.

37. (Previously Presented) A method for making an antimicrobial sulfonamide derivative comprising sulfonylating a core antibiotic or core cyclic peptide with a lipophilic sulfonyl derivative, thereby providing an antimicrobial sulfonamide derivative.

38. (Currently Amended) The method of Claim 37 in which the lipophilic sulfonyl derivative is ~~a~~ an activated lipophilic sulfonyl ester or a lipophilic sulfonyl halide.

39. (Original) The method of Claim 38 in which the activated lipophilic sulfonyl ester is a lipophilic hydroxybenzotriazole ester.

40. (Previously Presented) The method of Claim 38 in which the lipophilic sulfonyl halide is a lipophilic sulfonyl chloride.

41. (Currently Amended) A method for making an antimicrobial sulfonamide derivative comprising:

sulfonylating a linker with a lipophilic sulfonyl compound, thereby providing a lipophilic sulfonamide linker; and

covalently attaching the lipophilic sulfonamide linker to a core antibiotic or core cyclic peptide wherein said core cyclic peptide or core antibiotic is ~~not of polymyxin of~~ an acidic lipopeptide antibiotic, thereby yielding an antimicrobial sulfonamide derivative.

42. (Previously Presented) A method for making an antimicrobial sulfonamide derivative comprising:

covalently attaching a linker to a core antibiotic or core cyclic peptide, thereby providing an linker core antibiotic or linker core cyclic peptide; and

sulfonylating the linker core antibiotic or linker core cyclic peptide with a lipophilic sulfonyl derivative, thereby yielding an antimicrobial sulfonamide derivative.

43. (New) A method for treating or preventing a microbial infection, said method comprising the step of administering to a subject a therapeutically effective amount of an antimicrobial sulfonamide derivative according to any one of Claims 1 to 5.

44. (New) The method of Claim 43 in which the core cyclic peptide is aspartocin.

45. (New) The method of Claim 43 in which the core antibiotic is aspartocin.

46. (New) A method of inhibiting microbial growth, said method comprising the step of administering to a microbe an antimicrobially effective amount of an antimicrobial sulfonamide derivative according to any one of Claims 1 to 5.

47. (New) The method of Claim 46 in which the core cyclic peptide is aspartocin.

48. (New) The method of Claim 46 in which the core antibiotic is aspartocin.